

SEQUENCE LISTING

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<110>

Mount David B.
Delpire, Eric
Gamba, Gerardo
Alfred L. George, Jr.

<120> PURIFIED AND ISOLATED POTASSIUM-CHLORIDE COTRANSPORTER NUCLEIC ACIDS AND POLYPEPTIDES AND THERAPEUTIC AND SCREENING METHODS USING SAME

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<130> Attorney Docket No. 1242-26-2
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pur pi

 $\mathcal{A}^{\mathsf{l}}$ 

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July Const

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July 180

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Cys Thr Cys Thr Met Leu Thr Ala Ile Ser Met Ser Ala Ile Ala Thr 165 170 175

Asn Gly Val Val Pro Ala Gly Gly Ser Tyr Tyr Met Ile Ser Arg Ser 180 185 190

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Leu Thr Tyr Ile Ser Pro Gly Ala Ala Ile Phe Gln Ala Glu Ala Ala 225 230 235 240

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Met Phe Leu Tyr His Leu Arg Ile Ser Ala Glu Val Glu Val Val Glu 900 905 910

Met Val Glu Asn Asp Ile Ser Ala Phe Thr Tyr Glu Arg Thr Leu Met 915 920 925

Met Glu Gln Arg Ser Gln Met Leu Lys Gln Met Gln Leu Ser Lys Asn 930 935 940

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Pro	Asp	Xaa 35	Ser	Ser	Arg	Ser	Ser 40	Ser	Arg	Val	Arg	Phe 45	Ser	Ser	Arg
Glu	Ser 50	Val	Pro	Glu	Thr	Ser 55	Arg	Ser	Glu	Pro	Met 60	Ser	Glu	Met	Ser
Gly 65	Ala	Thr	Thr	Ser	Leu 70	Ala	Thr	Val	Ala	Leu 75	Asp	Pro	Pro	Ser	Asp 80
Arg	Thr	Ser	His	Pro 85	Gln	Asp	Val	Ile	Glu 90	Asp	Asp	Gly	His	Lys 95	Lys
Ala	Arg	Asn	Ala 100	Tyr	Leu	Asn	Asn	Ser 105	Asn	Tyr	Glu	Glu	Gly 110	Asp	Glu
Tyr	Phe	Asp 115	Lys	Asn	Leu	Ala	Leu 120	Phe	Glu	Glu	Glu	Met 125	Asp	Thr	Arg
Pro	Lys 130	Val	Ser	Ser	Leu	Leu 135	Asn	Arg	Met	Ala	Asn 140	Tyr	Thr	Asn	Leu
Thr 145	Gln	Gly	Ala	Lys	Glu 150	His	Glu	Glu	Ala	Glu 155	Asn	Ile	Thr	Glu	Gly 160
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Val Leu Ile Cys Cys Cys Cys Thr Met Leu Thr Ala Ile Ser Met Ser

170

220

Ala Ile Ala Thr Asn Gly Val Val Pro Ala Gly Gly Ser Tyr Phe Met 225 230 235 240

Ile Ser Arg Ala Leu Gly Pro Glu Phe Gly Gly Ala Val Gly Leu Cys 245 Phe Tyr Leu Gly Thr Thr Phe Ala Ala Ala Met Tyr Ile Leu Gly Ala 265 Ile Glu Ile Phe Leu Val Tyr Ile Val Pro Arg Ala Ala Ile Phe His 275 Ser Asp Asp Ala Leu Lys Glu Ser Ala Ala Met Leu Asn Asn Met Arq Val Tyr Gly Thr Ala Phe Leu Val Leu Met Val Leu Val Val Phe Ile 310 315 Gly Val Arg Tyr Val Asn Lys Phe Ala Ser Leu Phe Leu Ala Cys Val 325 Ile Val Ser Ile Leu Ala Ile Tyr Ala Gly Ala Ile Lys Ser Ser Phe Ala Pro Pro His Phe Pro Val Cys Met Leu Gly Asn Arg Thr Leu Ser 360 Ser Arg His Ile Asp Val Cys Ser Lys Thr Lys Glu Ile Asn Asn Met 375 Thr Val Pro Ser Lys Leu Trp Gly Phe Phe Cys Asn Ser Ser Gln Phe 390 395 Phe Asn Ala Thr Cys Asp Glu Tyr Phe Val His Asn Asn Val Thr Ser 410 405 Ile Gln Gly Ile Pro'Gly Leu Ala Ser Gly Ile Ile Thr Glu Asn Leu 425 Trp Ser Asn Tyr Leu Pro Lys Gly Glu Ile Ile Glu Lys Pro Ser Ala 440 Lys Ser Ser Asp Val Leu Gly Ser Leu Asn His Glu Tyr Val Leu Val 450 455 Asp Ile Thr Thr Ser Phe Thr Xaa Leu Val Gly Ile Phe Phe Pro Ser 470 Val Thr Gly Ile Met Ala Gly Ser Asn Arg Ser Gly Asp Leu Lys Asp Ala Gln Lys Ser Ile Pro Ile Gly Thr Ile Leu Ala Ile Leu Thr Thr 500 505 Ser Phe Val Tyr Leu Ser Asn Val Val Leu Phe Gly Ala Cys Ile Glu 520 Gly Val Val Leu Arg Asp Lys Phe Gly Asp Ala Val Lys Gly Asn Leu 530 535 540

Val Val Gly Thr Leu Ser Trp Pro Ser Pro Trp Val Ile Val Ile Gly 550 Ser Phe Phe Ser Thr Cys Gly Ala Gly Leu Gln Ser Leu Thr Gly Ala Pro Arq Leu Leu Gln Ala Ile Ala Lys Asp Asn Ile Ile Pro Phe Leu Arg Val Phe Gly His Ser Lys Ala Asn Gly Glu Pro Thr Trp Ala Leu Leu Leu Thr Ala Ala Ile Ala Glu Leu Gly Ile Leu Ile Ala Ser Leu 615 Asp Leu Val Ala Pro Ile Leu Ser Met Phe Phe Leu Met Cys Tyr Leu 630 635 Phe Val Asn Leu Ala Cys Ala Leu Gln Thr Leu Leu Arg Thr Pro Asn 650 Trp Arg Pro Arg Phe Arg Tyr Tyr His Trp Ala Leu Ser Phe Met Gly 660 665 Met Ser Ile Cys Leu Ala Leu Met Phe Ile Ser Ser Trp Tyr Tyr Ala 680 Ile Val Ala Met Val Ile Ala Gly Met Ile Tyr Lys Tyr Ile Glu Tyr 695 Gln Gly Ala Glu Lys Glu Trp Gly Asp Gly Ile Arg Gly Leu Ser Leu Ser Ala Ala Arg Phe Ala Leu Leu Arg Leu Glu Glu Gly Pro Pro His Thr Lys Asn Trp Arg Pro Gln Leu Leu Val Leu Leu Lys Leu Asp Glu 740 745 Asp Leu His Val Lys His Pro Arg Leu Leu Thr Phe Ala Ser Gln Leu Lys Ala Gly Lys Gly Leu Thr Ile Val Gly Ser Val Ile Val Gly Asn 775 Phe Leu Glu Asn Tyr Gly Glu Ala Leu Ala Ala Glu Gln Thr Ile Lys 785 790 His Leu Met Glu Ala Glu Lys Val Lys Gly Phe Cys Gln Leu Val Val 810 805 Ala Ala Lys Leu Arg Glu Gly Ile Ser His Leu Ile Gln Ser Cys Gly 825 Leu Gly Gly Met Lys His Asn Thr Val Val Met Gly Trp Pro Asn Gly 840 835

Trp Arg Gln Ser Glu Asp Ala Arg Ala Trp Lys Thr Phe Ile Gly Thr 850 860

Val Arg Val Thr Thr Ala Ala His Leu Ala Leu Leu Val Ala Lys Asn 865 870 875 880

Ile Ser Phe Phe Pro Ser Asn Val Glu Gln Phe Ser Glu Gly Asn Ile 885 890 895

Asp Val Trp Trp Ile Val His Asp Gly Gly Met Leu Met Leu Leu Pro 900 905 910

Phe Leu Leu Lys Gln His Lys Val Trp Arg Lys Cys Ser Ile Arg Ile 915 920 925

Phe Thr Val Ala Gln Leu Glu Asp Asn Ser Ile Gln Met Lys Lys Asp 930 935 940

Leu Ala Thr Phe Leu Tyr His Leu Arg Ile Glu Ala Glu Val Glu Val 945 950 955 960

Val Glu Met His Asp Ser Asp Ile Ser Ala Tyr Thr Tyr Glu Arg Thr
965 970 975

Leu Met Met Glu Gln Arg Ser Gln Met Leu Arg His Met Arg Leu Ser 980 985 990

Lys Thr Glu Arg Asp Arg Glu Ala Gln Leu Val Lys Asp Arg Asn Ser 995 1000 1005

Met Leu Arg Leu Thr Ser Ile Gly Ser Asp Glu Asp Glu Glu Thr Glu 1010 1015 1020

Thr Tyr Gln Glu Lys Val His Met Thr Trp Thr Lys Asp Lys Tyr Met 1025 1030 1035 1040

Ala Ser Arg Gly Gln Lys Ala Lys Ser Met Glu Gly Phe Gln Asp Leu 1045 1050 1055

Leu Asn Met Arg Pro Asp Gln Ser Asn Val Arg Arg Met His Thr Ala 1060 1065 1070

Val Lys Xaa Asn Glu Val Ile Val Asn Lys Ser His Glu Ala Lys Leu 1075 1080 1085

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Met Val Thr Pro Thr Lys Ile Asp Asp Ile Pro Gly Leu Ser Asp Thr
age eeg gae etc age tet egg tet agt tet ega gta aga tit age tee
Ser Pro Asp Leu Ser Ser Arg Ser Ser Ser Arg Val Arg Phe Ser Ser
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Arg Glu Ser Val Pro Glu Thr Ser Arg Ser Glu Pro Met Ser Glu Leu
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                                                                   240
Ser Gly Ala Thr Thr Ser Leu Ala Thr Val Ala Leu Asp Pro Ser Ser
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Asp Arg Thr Ser Asn Pro Gln Asp Val Thr Glu Asp Asp Gly His Lys
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Lys Ala Arg Asn Ala Tyr Xaa Asn Asn Ser Asn Tyr Glu Glu Gly Asp
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gaa tat ttt gat aaa aat ttg gca ctc ttt gag gaa gaa atg gac acc
                                                                   384
Glu Tyr Phe Asp Lys Asn Leu Ala Leu Phe Glu Glu Glu Met Asp Thr
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                                120
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				gaa Glu 150										480
				aag Lys										528
				cag Gln										576
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				tgc Cys										672
				gga Gly 230										720
				ggc Gly										768
				aca Thr										816
				gta Val										864
				aag Lys									_	912
_	_		-	ttc Phe 310	_	_		_	_	-		_		960
				aat Asn										1008
_		 _	_	gct Ala			_		_		_			1056

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		gta Val														1200
		aat Asn														1248
		caa Gln														1296
		agt Ser														1344
		tca Ser 450														1392
		atc Ile														1440
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		cag Gln														1536
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		gta Val														1680

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						ata Ile									1872
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				_		cgc Arg				 _				_	2016
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•

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•						aaa Lys											2496
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						act Thr											2640
						ccc Pro 885											2688
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Glu Ser Val Pro Glu Thr Ser Arg Ser Glu Pro Met Ser Glu Leu Ser 50 55 60

Gly Ala Thr Thr Ser Leu Ala Thr Val Ala Leu Asp Pro Ser Ser Asp 65 70 75 80

Arg Thr Ser Asn Pro Gln Asp Val Thr Glu Asp Asp Gly His Lys Lys
85 90 95

Ala Arg Asn Ala Tyr Xaa Asn Asn Ser Asn Tyr Glu Glu Gly Asp Glu
100 105 110

Tyr Phe Asp Lys Asn Leu Ala Leu Phe Glu Glu Met Asp Thr Arg
115 120 125

Pro Lys Val Ser Ser Leu Leu Asn Arg Met Ala Asn Tyr Thr Asn Leu 130 135 140

Thr Gln Gly Ala Lys Glu His Glu Glu Ala Glu Asn Ile Thr Glu Gly 145 150 155 160

Lys Lys Lys Pro Thr Lys Ser Pro Gln Met Gly Thr Phe Met Gly Val 170 165 Tyr Leu Pro Cys Leu Gln Asn Ile Phe Gly Val Ile Leu Phe Leu Arg 185 Leu Thr Trp Val Val Gly Thr Ala Gly Ile Leu Gln Ala Phe Ala Ile 200 195 Val Leu Ile Cys Cys Cys Cys Thr Met Leu Thr Ala Ile Ser Met Ser Ala Ile Ala Thr Asn Gly Val Val Pro Ala Gly Gly Ser Tyr Phe Met 230 235 Ile Ser Arg Ala Leu Gly Pro Glu Phe Gly Gly Ala Val Gly Leu Cys 245 Phe Tyr Leu Gly Thr Thr Phe Ala Ala Met Tyr Ile Leu Gly Ala 265 Ile Glu Ile Phe Leu Val Tyr Ile Val Pro Arg Ala Ala Ile Phe Arg Ser Asp Asp Ala Leu Lys Glu Ser Ala Ala Met Leu Asn Asn Met Arg 290 295 Val Tyr Gly Thr Ala Phe Leu Val Leu Met Val Leu Val Val Phe Ile Gly Val Arg Tyr Val Asn Lys Phe Ala Ser Leu Phe Leu Ala Cys Val Ile Val Ser Ile Leu Ala Ile Tyr Ala Gly Ala Ile Lys Ser Ser Phe 340 345 Ala Pro Pro His Phe Pro Val Cys Met Leu Gly Asn Arg Thr Leu Ser 360 Ser Arg His Leu Asp Ile Cys Ser Lys Thr Lys Glu Val Asp Asn Met 375 370 Thr Val Pro Ser Lys Leu Trp Gly Phe Phe Cys Asn Ser Ser Gln Phe 390 395 Phe Asn Ala Thr Cys Asp Glu Tyr Phe Val His Asn Asn Val Ile Ser 405 410 Ile Gln Gly Ile Pro Gly Leu Ala Ser Gly Ile Ile Thr Glu Asn Leu Trp Ser Asn Tyr Leu Pro Lys Gly Glu Ile Ile Glu Lys Pro Ser Ala 440 Lys Ser Ser Asp Val Leu Gly Asn Leu Asn His Glu Tyr Val Leu Ala 450 460

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Lys Ala Gly Lys Gly Leu Thr Ile Val Gly Ser Val Ile Val Gly Asn 770 780

Phe Leu Glu Asn Tyr Gly Asp Ala Leu Ala Ala Glu Gln Thr Ile Lys 785 790 795 800

His Leu Met Glu Ala Glu Lys Val Lys Gly Phe Cys Gln Leu Val Val 805 810 815

Ala Ala Lys Leu Lys Glu Gly Ile Ser His Leu Ile Gln Ser Cys Gly
820 825 830

Leu Gly Gly Met Lys His Asn Thr Val Val Met Gly Trp Pro Asn Gly 835 840 845

Trp Arg Gln Ser Glu Asp Ala Arg Ala Trp Lys Thr Phe Ile Gly Thr 850 855 860

Val Arg Val Thr Thr Ala Ala His Leu Ala Leu Leu Val Ala Lys Asn 865 870 875 880

Val Ser Phe Phe Pro Ser Asn Val Glu Gln Phe Ser Glu Gly Asn Ile 885 890 895

Asp Val Arg Trp Ile Val His Asp Gly Gly Met Leu Met Leu Pro 900 905 910

Phe Leu Leu Lys Gln His Lys Val Trp Arg Lys Cys Ser Ile Arg Ile 915 920 925

Phe Thr Val Ala Gln Leu Glu Asp Asn Ser Ile Gln Met Lys Lys Asp 930 935 940

Leu Ala Thr Phe Leu Tyr His Leu Arg Ile Glu Ala Glu Val Glu Val 945 950 955 960

Val Glu Met His Asp Ser Asp Ile Ser Ala Tyr Thr Tyr Glu Arg Thr 965 970 975

Leu Met Met Glu Gln Arg Ser Gln Met Leu Arg His Met Arg Leu Ser 980 985 990

Lys Thr Glu Arg Asp Arg Glu Ala Gln Leu Val Lys Asp Arg Asn Ser 995 1000 1005

Met Leu Arg Leu Thr Ser Ile Gly Ser Asp Glu Asp Glu Glu Thr Glu 1010 1015 1020

Thr Tyr Gln Glu Lys Val His Met Thr Trp Thr Lys Asp Lys Tyr Met 1025 1030 1035 1040

Ala Ser Arg Gly Gln Lys Val Lys Ser Met Glu Gly Phe Gln Asp Leu 1045 1050 1055

Leu Asn Met Arg Pro Asp Gln Ser Asn Val Arg Arg Met His Thr Ala 1060 1065 1070

Val Lys Leu Asn Glu Val Ile Val Asn Lys Ser His Glu Ala Lys Leu 1075 1080 Val Leu Leu Asn Met Pro Gly Pro Pro Arg Asn Pro Glu Gly Asp Glu 1095 Asn Tyr Met Glu Phe Leu Glu Val Xaa Thr Glu Gly Leu Glu Arg Val 1105 1110 1115 Leu Leu Val Arg Gly Gly Gly Ser Glu Val Ile Thr Ile Tyr Ser 1125 1130 <210> 7 <211> 6120 <212> DNA <213> mouse <220> <221> CDS <222> (4)..(3453) <220> <221> misc feature <222> (106) <223> n=c or a, Xaa=Leu or Ile <220> <221> misc\_feature <222> (925) <223> n=c or a, Xaa=Leu or Ile <220> <221> misc feature <222> (2545) <223> n=c or a, Xaa=Leu or Ile aaa atg cat cca cca gaa gcc acc acc aag atg tcc tca gtt cgg ttc Met His Pro Pro Glu Ala Thr Thr Lys Met Ser Ser Val Arg Phe atg gtg aca cca act aag att gat gac att cca ggt ttg tca gac acc Met Val Thr Pro Thr Lys Ile Asp Asp Ile Pro Gly Leu Ser Asp Thr age eeg gae nte age tet egg tet agt tet ega gta aga tit age tee 144 Ser Pro Asp Xaa Ser Ser Arg Ser Ser Ser Arg Val Arg Phe Ser Ser cga gaa agt gtg cca gaa aca agc cgt agt gag cct atg agc gaa ctg 192 Arg Glu Ser Val Pro Glu Thr Ser Arg Ser Glu Pro Met Ser Glu Leu 50 tot ggg gct act act tot otg gca act gtt gcc cta gat cot toc agt Ser Gly Ala Thr Thr Ser Leu Ala Thr Val Ala Leu Asp Pro Ser Ser 65

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			att Ile 645								1968
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Glu Ser Val Pro Glu Thr Ser Arg Ser Glu Pro Met Ser Glu Leu Ser 50 55 60

Gly Ala Thr Thr Ser Leu Ala Thr Val Ala Leu Asp Pro Ser Ser Asp
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Arg Thr Ser Asn Pro Gln Asp Val Thr Glu Asp Pro Ser Gln Asn Ser Ile Thr Gly Glu His Ser Gln Leu Leu Asp Asp Gly His Lys Lys Ala Arg Asn Ala Tyr Leu Asn Asn Ser Asn Tyr Glu Glu Gly Asp Glu Tyr 120 Phe Asp Lys Asn Leu Ala Leu Phe Glu Glu Glu Met Asp Thr Arg Pro Lys Val Ser Ser Leu Leu Asn Arg Met Ala Asn Tyr Thr Asn Leu Thr Gln Gly Ala Lys Glu His Glu Glu Ala Glu Asn Ile Thr Glu Gly Lys 170 Lys Lys Pro Thr Lys Ser Pro Gln Met Gly Thr Phe Met Gly Val Tyr 190 180 185 Leu Pro Cys Leu Gln Asn Ile Phe Gly Val Ile Leu Phe Leu Arg Leu Thr Trp Val Val Gly Thr Ala Gly Ile Leu Gln Ala Phe Ala Ile Val 215 Leu Ile Cys Cys Cys Cys Thr Met Leu Thr Ala Ile Ser Met Ser Ala 225 Ile Ala Thr Asn Gly Val Val Pro Ala Gly Gly Ser Tyr Phe Met Ile Ser Arg Ala Leu Gly Pro Glu Phe Gly Gly Ala Val Gly Leu Cys Phe Tyr Leu Gly Thr Thr Phe Ala Ala Ala Met Tyr Ile Leu Gly Ala Ile 275 Glu Ile Phe Leu Val Tyr Ile Val Pro Arg Ala Ala Ile Phe Arg Ser 295 Asp Asp Ala Leu Lys Glu Ser Ala Ala Met Leu Asn Asn Met Arg Val 305 Tyr Gly Thr Ala Phe Leu Val Leu Met Val Leu Val Val Phe Ile Gly 325 330 Val Arg Tyr Val Asn Lys Phe Ala Ser Leu Phe Leu Ala Cys Val Ile 345 Val Ser Ile Leu Ala Ile Tyr Ala Gly Ala Ile Lys Ser Ser Phe Ala 355 Pro Pro His Phe Pro Val Cys Met Leu Gly Asn Arg Thr Leu Ser Ser 375 380 370

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Ser Ile Cys Leu Ala Leu Met Phe Ile Ser Ser Trp Tyr Tyr Ala Ile 690 695 700

Val Ala Met Val Ile Ala Gly Met Ile Tyr Lys Tyr Ile Glu Tyr Gln 705 710 715 720

Gly Ala Glu Lys Glu Trp Gly Asp Gly Ile Arg Gly Leu Ser Leu Ser 725 730 735

Ala Ala Arg Phe Ala Leu Leu Arg Leu Glu Glu Gly Pro Pro His Thr
740 745 750

Lys Asn Trp Arg Pro Gln Leu Leu Val Leu Leu Lys Leu Asp Glu Asp 755 760 765

Leu His Val Lys His Pro Arg Leu Leu Thr Phe Ala Ser Gln Leu Lys
770 780

Ala Gly Lys Gly Leu Thr Ile Val Gly Ser Val Ile Val Gly Asn Phe
785 790 795 800

Leu Glu Asn Tyr Gly Asp Ala Leu Ala Ala Glu Gln Thr Ile Lys His 805 810 815

Leu Met Glu Ala Glu Lys Val Lys Gly Phe Cys Gln Leu Val Val Ala 820 825 830

Ala Lys Leu Lys Glu Gly Ile Ser His Leu Ile Gln Ser Cys Gly Leu 835 840 845

Gly Gly Met Lys His Asn Thr Val Val Met Gly Trp Pro Asn Gly Trp 850 855 860

Arg Gln Ser Glu Asp Ala Arg Ala Trp Lys Thr Phe Ile Gly Thr Val 865 870 875 880

Arg Val Thr Thr Ala Ala His Leu Ala Leu Leu Val Ala Lys Asn Val 885 890 895

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Thr Val Ala Gln Leu Glu Asp Asn Ser Ile Gln Met Lys Lys Asp Leu 945 950 955 960

Ala Thr Phe Leu Tyr His Leu Arg Ile Glu Ala Glu Val Glu Val Val 965 970 975

Glu Met His Asp Ser Asp Ile Ser Ala Tyr Thr Tyr Glu Arg Thr Leu 980 985 990

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										ccc Pro					496
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Cys Asp Glu Tyr Phe Val His Asn Asn Val Ile Ser Ile Gln Gly Ile 370 375 380

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Leu Pro Lys Gly Glu Ile Ile Glu Lys Pro Ser Ala Lys Ser Ser Asp 405 410 415

Val Leu Gly Asn Leu Asn His Glu Tyr Val Leu Ala Asp Ile Thr Thr 420 425 430

Ser Phe Thr Leu Leu Val Gly Ile Phe Phe Pro Ser Val Thr Gly Ile 435 440 445

Met Ala Gly Ser Asn Arg Ser Gly Asp Leu Lys Asp Ala Gln Lys Ser 450 455 460

Ile Pro Ile Gly Thr Ile Leu Ala Ile Leu Thr Thr Ser Phe Val Tyr 465 470 475 480

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ccc gac aaa gtg Pro Asp Lys Val 975		r Trp Thr	Lys Glu L			3039
aaa cac agg aac Lys His Arg Asr 990	aag gac ac Lys Asp Th 995	t ggc cca r Gly Pro	tca ggc t Ser Gly P 1000	tc aaa gac Phe Lys Asp	ctc ttc Leu Phe 1005	3087
agc cta aag ccg Ser Leu Lys Pro		r Asn Val				3135
aag ntc aac ggo Lys Xaa Asn Gly 102	Val Val Le		Ser Gln A		Leu Val	3183
ctg ctg aat atg Leu Leu Asn Met 1040						3231
tac atg gag tto Tyr Met Glu Phe 1055		l Leu Thr	Glu Gly I			3279
ctg gtc agg ggt Leu Val Arg Gly 1070	ggt ggc cg Gly Gly Ar 1075	a gaa gtc g Glu Val	atc acc a Ile Thr I 1080	atc tac tcc [le Tyr Ser		3321
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Asn Ser Pro Phe Ile Asn Asn Val Glu Val Glu Arg Glu Ser Tyr Phe
    50
Glu Gly Lys Asn Met Ala Xaa Phe Glu Glu Glu Met Asp Ser Asn Pro
Met Val Ser Ser Leu Leu Asn Lys Leu Ala Asn Tyr Thr Asn Leu Ser
Gln Gly Val Val Glu His Glu Glu Asp Glu Asp Ser Arg Arg Glu
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Leu Gln Asn Ile Leu Gly Val Ile Leu Phe Leu Arg Leu Thr Trp Ile
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Val Gly Ala Ala Gly Val Met Glu Ser Phe Xaa Ile Val Ala Met Cys
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Cys Thr Cys Thr Met Leu Thr Ala Ile Ser Met Ser Ala Ile Ala Thr
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Asn Gly Val Val Pro Ala Gly Gly Ser Tyr Tyr Met Ile Ser Arg Ser
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Gly Met Leu Ala Trp Pro Ser Pro Trp Val Ile Val Ile Gly Ser Phe 500 Trp Sos Leu Thr Gly Ala Pro Arg

515 520 525

Leu Leu Gln Ala Ile Ala Arg Asp Gly Ile Ile Pro Phe Leu Gln Val 530 540

Phe Gly His Gly Lys Ala Asn Gly Glu Pro Thr Trp Ala Leu Leu Leu 545 550 555 560

Thr Ala Leu Ile Cys Glu Thr Gly Ile Leu Ile Ala Ser Leu Asp Ser 565 570 575

Val Ala Pro Ile Leu Ser Met Phe Phe Xaa Met Cys Tyr Met Phe Val 580 585 590

Asn Leu Ala Cys Ala Val Gln Thr Leu Leu Arg Thr Pro Asn Trp Arg 595 600 605

Pro Arg Phe Lys Phe Tyr His Trp Thr Leu Ser Phe Leu Gly Met Ser 610 620

Leu Cys Leu Ala Leu Met Phe Ile Cys Ser Trp Tyr Tyr Ala Xaa Phe 625 630 635 640

Ala Met Leu Ile Ala Gly Cys Ile Tyr Lys Tyr Ile Glu Tyr Arg Gly 645 650 655

Ala Glu Lys Glu Trp Gly Asp Gly Ile Arg Gly Leu Ser Leu Asn Ala 660 665 670

Ala Arg Tyr Ala Leu Leu Arg Val Glu His Gly Pro Pro His Thr Lys
675 680 685

Asn Trp Arg Pro Gln Val Leu Val Met Leu Asn Leu Asp Ser Glu Gln 690 695 700

Cys Val Lys His Pro Arg Leu Leu Ser Phe Thr Ser Gln Leu Lys Ala 705 710 715 720

Gly Lys Gly Leu Thr Ile Val Gly Ser Val Leu Glu Gly Thr Tyr Leu
725 730 735

Asp Lys His Val Glu Ala Gln Arg Ala Glu Glu Asn Ile Arg Ser Leu
740 745 750

Met Ser Ala Glu Lys Thr Lys Gly Phe Cys Gln Leu Val Val Ser Ser 755 760 765

Asn Leu Arg Asp Gly Ala Ser His Leu Ile Gln Ser Ala Gly Leu Gly
770 775 780

Gly Met Lys His Asn Thr Val Leu Met Ala Trp Pro Glu Ala Trp Lys
785 790 795 800

Glu Ala Asp Asn Pro Phe Ser Trp Lys Asn Phe Val Asp Thr Val Arg 805 810 815

Asp Thr Thr Ala Ala His Gln Ala Leu Leu Val Ala Lys Asn Ile Asp 820 825 830

Leu Phe Pro Gln Asn Gln Glu Arg Phe Ser Asp Gly Asn Ile Asp Val 835 840 845

Trp Trp Ile Val His Asp Gly Gly Met Leu Met Leu Leu Pro Phe Leu 850 855 860

Leu Arg Gln His Lys Val Trp Arg Lys Cys Arg Met Arg Ile Phe Thr 865 870 875 880

Val Ala Gln Val Asp Asp Asn Ser Ile Gln Met Lys Lys Asp Leu Gln 885 890 895

Met Phe Leu Tyr His Leu Arg Ile Ser Ala Glu Val Glu Val Glu 900 905 910

Met Val Glu Asn Asp Ile Ser Ala Phe Thr Tyr Glu Lys Thr Leu Met 915 920 925

Met Glu Gln Arg Ser Gln Met Leu Lys Gln Met Gln Leu Ser Lys Asn 930 935 940

Glu Arg Glu Arg Glu Ala Gln Leu Ile His Asp Arg Asn Thr Ala Ser 945 950 955 960

His Thr Thr Ala Thr Ala Arg Thr Gln Ala Pro Pro Thr Pro Asp Lys
965 970 975

Val Gln Met Thr Trp Thr Lys Glu Lys Leu Ile Ala Glu Lys His Arg 980 985 990

Asn Lys Asp Thr Gly Pro Ser Gly Phe Lys Asp Leu Phe Ser Leu Lys 995 1000 1005

Pro Asp Gln Ser Asn Val Arg Arg Met His Thr Ala Val Lys Xaa Asn 1010 1015 1020

Gly Val Val Leu Asn Lys Ser Gln Asp Ala Gln Leu Val Leu Leu Asn 1025 1030 1035 1040

Met Pro Gly Pro Pro Lys Ser Arg Gln Gly Asp Glu Asn Tyr Met Glu 1045 1050 1055

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tttgcgaggt tccccccact ttttttgttc ttaaaaagaa caaa atg cat cct cca
                                                 Met His Pro Pro
                                                                   224
gaa acc acc acc aag atg gct tca gtt cgg ttc atg gtg aca ccg aca
Glu Thr Thr Lys Met Ala Ser Val Arg Phe Met Val Thr Pro Thr
aag atc gat gac att cca ggt ttg tca gac acc agt ccg gac ntc agc
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Lys Ile Asp Asp Ile Pro Gly Leu Ser Asp Thr Ser Pro Asp Xaa Ser
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tot cga tot agt toc cga gta aga ttt agc toc cgg gaa agc gtg cot
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Ser Arg Ser Ser Ser Arg Val Arg Phe Ser Ser Arg Glu Ser Val Pro
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gaa aca agc cgg agt gag cct atg agt gag atg tct ggg gcc acc act
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Glu Thr Ser Arg Ser Glu Pro Met Ser Glu Met Ser Gly Ala Thr Thr
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tcg ctg gca act gtt gca ctg gat cca ccc agt gac cgg act tct cac
Ser Leu Ala Thr Val Ala Leu Asp Pro Pro Ser Asp Arg Thr Ser His
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                         75
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Pro Gln Asp Val Ile Glu Asp Leu Ser Gln Asn Ser Ile Thr Gly Glu
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His Ser Gln Leu Leu Asp Asp Gly His Lys Lys Ala Arg Asn Ala Tyr
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Cys	Cys 230	Thr	Met	Leu	Thr	gct Ala 235	Ile	Ser	Met	Ser	Ala 240	Ile	Ala	Thr	Asn	896
Gly 245	Val	Val	Pro	Ala	Gly 250	ggc Gly	Ser	Tyr	Phe	Met 255	Ile	Ser	Arg	Ala	Leu 260	944
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						tta Leu										1184

						ttc Phe						ttg Leu	1232
						atc Ile							1280
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				ata Ile							2096
				ttt Phe 650							2144
				tta Leu							2192
				gcc Ala							2240
				tct Ser							2288
				tac Tyr							2336
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				gag Glu							2432
				tta Leu							2480
				acc Thr							2528

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					ctc Leu								2720
		_			atg Met							-	2768
-	_	_	-		aag Lys				_	_			2816
					ctg Leu 890								2864
					ttt Phe								2912
					atg Met								2960
					aag Lys								3008
					atc Ile								3056
					gag Glu 970								3104
					tat Tyr								3152
				Leu	cgg Arg			Leu			Arg		3200

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agc att ggc tct gat gag gac gaa gag aca gaa acc tat cag gag aag 3296 Ser Ile Gly Ser Asp Glu Asp Glu Glu Thr Glu Thr Tyr Gln Glu Lys 1030 1035 1040
gtg cac atg act tgg aca aaa gac aag tac atg gca tcc cgg gga caa 3344 Val His Met Thr Trp Thr Lys Asp Lys Tyr Met Ala Ser Arg Gly Gln 1045 1050 1055 1060
aaa gcg aag tca atg gaa gga ttc cag gac ctg ctt aac atg cgt ccg 3392 Lys Ala Lys Ser Met Glu Gly Phe Gln Asp Leu Leu Asn Met Arg Pro 1065 1070 1075
gac cag tcc aat gtg agg cgg atg cat aca gca gtg aaa ctc aac gag 3440 Asp Gln Ser Asn Val Arg Arg Met His Thr Ala Val Lys Leu Asn Glu 1080 1085 1090
gtt ata gtt aac aag tcc cat gaa gca aag ctg gtt tta ttg aat atg 3488 Val Ile Val Asn Lys Ser His Glu Ala Lys Leu Val Leu Leu Asn Met 1095 1100 1105
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Glu Ser Val Pro Glu Thr Ser Arg Ser Glu Pro Met Ser Glu Met Ser
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Arg Thr Ser His Pro Gln Asp Val Ile Glu Asp Leu Ser Gln Asn Ser
Ile Thr Gly Glu His Ser Gln Leu Leu Asp Asp Gly His Lys Lys Ala
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Arg Asn Ala Tyr Leu Asn Asn Ser Asn Tyr Glu Glu Gly Asp Glu Tyr
Phe Asp Lys Asn Leu Ala Leu Phe Glu Glu Glu Met Asp Thr Arg Pro
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Lys Val Ser Ser Leu Leu Asn Arg Met Ala Asn Tyr Thr Asn Leu Thr
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Gln Gly Ala Lys Glu His Glu Glu Ala Glu Asn Ile Thr Glu Gly Lys
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Ile Thr Thr Ser Phe Thr Leu Leu Val Gly Ile Phe Phe Pro Ser Val 490 485 Thr Gly Ile Met Ala Gly Ser Asn Arg Ser Gly Asp Leu Lys Asp Ala Gln Lys Ser Ile Pro Ile Gly Thr Ile Leu Ala Ile Leu Thr Thr Ser Phe Val Tyr Leu Ser Asn Val Val Leu Phe Gly Ala Cys Ile Glu Gly Val Val Leu Arg Asp Lys Phe Gly Asp Ala Val Lys Gly Asn Leu Val Val Gly Thr Leu Ser Trp Pro Ser Pro Trp Val Ile Val Ile Gly Ser 570 Phe Phe Ser Thr Cys Gly Ala Gly Leu Gln Ser Leu Thr Gly Ala Pro 585 Arg Leu Leu Gln Ala Ile Ala Lys Asp Asn Ile Ile Pro Phe Leu Arg 595 600 Val Phe Gly His Ser Lys Ala Asn Gly Glu Pro Thr Trp Ala Leu Leu 615 Leu Thr Ala Ala Ile Ala Glu Leu Gly Ile Leu Ile Ala Ser Leu Asp 630 635 Leu Val Ala Pro Ile Leu Ser Met Phe Phe Leu Met Cys Tyr Leu Phe 645 Val Asn Leu Ala Cys Ala Leu Gln Thr Leu Leu Arg Thr Pro Asn Trp 665 Arg Pro Arg Phe Arg Tyr Tyr His Trp Ala Leu Ser Phe Met Gly Met 680 Ser Ile Cys Leu Ala Leu Met Phe Ile Ser Ser Trp Tyr Tyr Ala Ile 695 Val Ala Met Val Ile Ala Gly Met Ile Tyr Lys Tyr Ile Glu Tyr Gln Gly Ala Glu Lys Glu Trp Gly Asp Gly Ile Arg Gly Leu Ser Leu Ser Ala Ala Arg Phe Ala Leu Leu Arg Leu Glu Glu Gly Pro Pro His Thr 745 Lys Asn Trp Arg Pro Gln Leu Leu Val Leu Leu Lys Leu Asp Glu Asp 760 Leu His Val Lys His Pro Arg Leu Leu Thr Phe Ala Ser Gln Leu Lys 770 775

Ala Gly Lys Gly Xaa Thr Ile Val Gly Ser Val Ile Val Gly Asn Phe 785 790 795 800

Leu Glu Asn Tyr Gly Glu Ala Leu Ala Ala Glu Gln Thr Ile Lys His 805 810 815

Leu Met Glu Ala Glu Lys Val Lys Gly Phe Cys Gln Leu Val Val Ala 820 825 830

Ala Lys Leu Arg Glu Gly Ile Ser His Leu Ile Gln Ser Cys Gly Leu 835 840 845

Gly Gly Met Lys His Asn Thr Val Val Met Gly Trp Pro Asn Gly Trp 850 855 860

Arg Gln Ser Glu Asp Ala Arg Ala Trp Lys Thr Phe Ile Gly Thr Val 865 870 875 880

Arg Val Thr Thr Ala Ala His Leu Ala Leu Leu Val Ala Lys Asn Ile 885 890 895

Ser Phe Phe Pro Ser Asn Val Glu Gln Phe Ser Glu Gly Asn Ile Asp 900 905 910

Val Trp Trp Ile Val His Asp Gly Gly Met Leu Met Leu Leu Pro Phe 915 920 925

Leu Leu Lys Gln His Lys Val Trp Arg Lys Cys Ser Ile Arg Ile Phe 930 935 940

Thr Val Ala Gln Leu Glu Asp Asn Ser Ile Gln Met Lys Lys Asp Leu 945 950 955 960

Ala Thr Phe Leu Tyr His Leu Arg Ile Glu Ala Glu Val Glu Val Val 965 970 975

Glu Met His Asp Ser Asp Ile Ser Ala Tyr Thr Tyr Glu Arg Thr Leu 980 985 990

Met Met Glu Gln Arg Ser Gln Met Leu Arg His Met Arg Leu Ser Lys 995 1000 1005

Thr Glu Arg Asp Arg Glu Ala Gln Leu Val Lys Asp Arg Asn Ser Met 1010 1015 1020

Leu Arg Leu Thr Ser Ile Gly Ser Asp Glu Asp Glu Glu Thr Glu Thr 1025 1030 1035 1040

Tyr Gln Glu Lys Val His Met Thr Trp Thr Lys Asp Lys Tyr Met Ala 1045 1050 1055

Ser Arg Gly Gln Lys Ala Lys Ser Met Glu Gly Phe Gln Asp Leu Leu 1060 1065 1070

Asn Met Arg Pro Asp Gln Ser Asn Val Arg Arg Met His Thr Ala Val 1075 1080 1085 Lys Leu Asn Glu Val Ile Val Asn Lys Ser His Glu Ala Lys Leu Val 1090 1095 1100

Leu Leu Asn Met Pro Gly Pro Pro Arg Asn Pro Glu Gly Asp Glu Asn 1105 1110 1115 1120

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Lys Val Lys Gly Phe Cys Gln Val Val Val Ala Gln Lys Leu Lys Glu 35 40 45

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Lys Val Trp Arg Lys Cys Lys Met Arg Ile Phe Thr Val Ala Gln Met 145 150 155 160

Glu Asp Asn Ser Ile Gln Met Lys Lys Asp Leu Ala Thr Phe Leu Tyr 165 170 175

His Leu Arg Ile Ala Ala Asp Val Glu Val Val Glu Met His Asp Ser 180 185 190

Asp Ile Ser Ala Tyr Thr Tyr Glu Arg Thr Leu Met Met Glu Gln Arg 195 200 205

Ser Gln Met Leu Arg Gln Met Arg Leu Ser Lys Thr Asp Arg Glu Arg 210 215 220

Glu Ala Gln Leu Val Lys Asp Arg Asn Ser Ile Leu Arg Leu Thr Ser 225 230 235 240

Val Gly Ser Asp Asp Asp Glu Asp Thr Glu Ala Ala Pro Glu Arg Val 245 250 255

His Met Thr Trp Thr Arg Asp Lys His His Ala Val Arg Val Ala Gln
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Ser Lys Pro Met Pro Ser Cys Gln Asp Leu Leu Asn Ile Arg Pro Asp 275 280 285

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